

SEQUENCE LISTING

<110> Penttila, Merja E.  
Ward, Michael  
Wang, Huaming  
Valkonen, Mari J.  
Saloheimo, Markku

<120> Increased Production of Secreted  
Proteins by Recombinant Eukaryotic Cells

<130> GC590-2

<140> US 09/816,227  
<141> 2001-03-23

<150> US 09/534,692  
<151> 2000-03-24

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<212> DNA  
<213> Trichoderma reesei

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<213> Trichoderma reesei

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Asp Ser Val Ala Asp Ile Asp Ser Arg Leu Ser Val Ile Pro Glu Ser						
50	55	60				
Gln Asp Ala Glu Asp Asp Glu Ser His Ser Thr Ser Ala Thr Ala Pro						
65	70	75	80			
Ser Thr Ser Glu Lys Lys Pro Val Lys Lys Arg Lys Ser Trp Gly Gln						
85	90	95				
Val Leu Pro Glu Pro Lys Thr Asn Leu Pro Pro Arg Lys Arg Ala Lys						
100	105	110				
Thr Glu Asp Glu Lys Glu Gln Arg Arg Val Glu Arg Val Leu Arg Asn						
115	120	125				
Arg Arg Ala Ala Gln Ser Ser Arg Glu Arg Lys Arg Leu Glu Val Glu						
130	135	140				
Ala Leu Glu Lys Arg Asn Lys Glu Leu Glu Thr Leu Leu Ile Asn Val						
145	150	155	160			
Gln Lys Thr Asn Leu Ile Leu Val Glu Glu Leu Asn Arg Phe Arg Arg						
165	170	175				
Ser Ser Gly Val Val Thr Arg Ser Ser Pro Leu Asp Ser Leu Gln						
180	185	190				
Asp Ser Ile Thr Leu Ser Gln Gln Leu Phe Gly Ser Arg Asp Gly Gln						
195	200	205				
Thr Met Ser Asn Pro Glu Gln Ser Leu Met Asp Gln Ile Met Arg Ser						
210	215	220				
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225	230	235	240			
Pro Ile Ser Asp Lys Glu Phe Gln Thr Lys Glu Glu Asp Glu Glu Gln						
245	250	255				
Ala Asp Glu Asp Glu Glu Met Glu Gln Thr Trp His Glu Thr Lys Glu						
260	265	270				
Ala Ala Ala Ala Lys Glu Lys Asn Ser Lys Gln Ser Arg Val Ser Thr						

275	280	285
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290	295	300
Pro Val Phe Ser Asp Asp Ala Gly Ala Asn Cys		Leu Gly Leu Asp Pro
305	310	315
Val His Gln Asp Asp Gly Pro Phe Ser Ile Gly		His Ser Phe Gly Leu
325	330	335
Ser Ala Ala Leu Asp Ala Asp Arg Tyr Leu Leu		Glu Ser Gln Leu Leu
340	345	350
Ala Ser Pro Asn Ala Ser Thr Val Asp Asp Asp		Tyr Leu Ala Gly Asp
355	360	365
Ser Ala Ala Cys Phe Thr Asn Pro Leu Pro Ser		Asp Tyr Asp Phe Asp
370	375	380
Ile Asn Asp Phe Leu Thr Asp Asp Ala Asn His		Ala Ala Tyr Asp Ile
385	390	395
Val Ala Ala Ser Asn Tyr Ala Ala Ala Asp Arg		Glu Leu Asp Leu Glu
405	410	415
Ile His Asp Pro Glu Asn Gln Ile Pro Ser Arg		His Ser Ile Gln Gln
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<211> 1615

<212> DNA

<213> Aspergillus nidulans

<400> 3

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tgagttgtat	gaaatcagca	gaccggttt	cgccagtgaa	aatggaggac	gctttcgcaa	240
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gtgtgatacc	tcaagagtca	actccttact	cctgctaata	actaccacag	aaaacgcgc	480
aagacagaag	atgagaaaaga	gcagcgccgg	attgagcag	ttcttcgcaa	ccgcgcagcc	540
gcacaaacct	ctcgcgagcg	caagagactt	gaaatggaga	atttagaaaag	cgagaagatt	600
gatatgaaac	aacaaaacca	gttccttctt	cagcgtctcg	cccagatgga	ggctgagaac	660
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<212> PRT  
<213> Aspergillus nidulans

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Lys Pro Glu Glu Lys Lys Pro Ala Lys Lys Arg Lys Ser Trp Gly Gln  
50 55 60  
Glu Leu Pro Val Pro Lys Thr Asn Leu Pro Pro Arg Lys Arg Ala Lys  
65 70 75 80  
Thr Glu Asp Glu Lys Glu Gln Arg Arg Ile Glu Arg Val Leu Arg Asn  
85 90 95  
Arg Ala Ala Ala Gln Thr Ser Arg Glu Arg Lys Arg Leu Glu Met Glu  
100 105 110  
Lys Leu Glu Ser Glu Lys Ile Asp Met Glu Gln Gln Asn Gln Phe Leu  
115 120 125  
Leu Gln Arg Leu Ala Gln Met Glu Ala Glu Asn Asn Arg Leu Ser Gln  
130 135 140  
Gln Val Ala Gln Leu Ser Ala Glu Val Arg Gly Ser Arg His Ser Thr  
145 150 155 160  
Pro Thr Ser Ser Ser Pro Ala Ser Val Ser Pro Thr Leu Thr Pro Thr  
165 170 175  
Leu Phe Lys Gln Glu Gly Asp Glu Val Pro Leu Asp Arg Ile Pro Phe  
180 185 190  
Pro Thr Pro Ser Val Thr Asp Tyr Ser Pro Thr Leu Lys Pro Ser Ser  
195 200 205  
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210 215 220  
Gly Leu Glu Gly Asp Glu Ser Ala Leu Thr Leu Phe Asp Leu Gly Ala  
225 230 235 240  
Ser Ile Lys His Glu Pro Thr His Asp Leu Thr Ala Pro Leu Ser Asp  
245 250 255  
Asp Asp Phe Arg Arg Leu Phe Asn Gly Asp Ser Ser Leu Glu Ser Asp  
260 265 270  
Ser Ser Leu Leu Glu Asp Gly Phe Ala Phe Asp Val Leu Asp Ser Gly  
275 280 285  
Asp Leu Ser Ala Phe Pro Phe Asp Ser Met Val Asp Phe Asp Thr Glu  
290 295 300  
Pro Val Thr Leu Glu Asp Leu Glu Gln Thr Asn Gly Leu Ser Asp Ser  
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<210> 5  
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<213> Trichoderma reesei

<400> 5

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35 40 45  
Asp Ser Val Ala Asp Ile Asp Ser Arg Leu Ser Val Ile Pro Glu Ser  
50 55 60  
Gln Asp Ala Glu Asp Asp Glu Ser His Ser Thr Ser Ala Thr Ala Pro  
65 70 75 80  
Ser Thr Ser Glu Lys Lys Pro Val Lys Lys Arg Lys Ser Trp Gly Gln  
85 90 95  
Val Leu Pro Glu Pro Lys Thr Asn Leu Pro Pro Arg Lys Arg Ala Lys  
100 105 110  
Thr Glu Asp Glu Lys Glu Gln Arg Arg Val Glu Arg Val Leu Arg Asn  
115 120 125  
Arg Arg Ala Ala Gln Ser Ser Arg Glu Arg Lys Arg Leu Glu Val Glu  
130 135 140  
Ala Leu Glu Lys Arg Asn Lys Glu Leu Glu Thr Leu Leu Ile Asn Val  
145 150 155 160  
Gln Lys Thr Asn Leu Ile Leu Val Glu Glu Leu Asn Arg Phe Arg Arg  
165 170 175  
Ser Ser Gly Val Val Thr Arg Ser Ser Pro Leu Asp Ser Leu Gln  
180 185 190  
Asp Ser Ile Thr Leu Ser Gln Gln Leu Phe Gly Ser Arg Asp Gly Gln  
195 200 205  
Thr Met Ser Asn Pro Glu Gln Ser Leu Met Asp Gln Ile Met Arg Ser  
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Ala Ala Asn Pro Thr Val Asn Pro Ala Ser Leu Ser Pro Ser Leu Pro  
225 230 235 240  
Pro Ile Ser Asp Lys Glu Phe Gln Thr Lys Glu Glu Asp Glu Glu Gln  
245 250 255  
Ala Asp Glu Asp Glu Glu Met Glu Gln Thr Trp His Glu Thr Lys Glu  
260 265 270  
Ala Ala Ala Ala Lys Glu Lys Asn Ser Lys Gln Ser Arg Val Ser Thr  
275 280 285  
Asp Ser Thr Gln Arg Pro Ala Val Ser Ile Gly Gly Asp Ala Ala Val  
290 295 300  
Pro Val Phe Ser Asp Asp Ala Gly Ala Asn Cys Leu Gly Leu Asp Pro  
305 310 315 320  
Val His Gln Asp Asp Gly Pro Phe Ser Ile Gly His Ser Phe Gly Leu  
325 330 335  
Ser Ala Ala Leu Asp Ala Asp Arg Tyr Leu Leu Glu Ser Gln Leu Leu  
340 345 350  
Ala Ser Pro Asn Ala Ser Thr Val Asp Asp Asp Tyr Leu Ala Gly Asp  
355 360 365  
Ser Ala Ala Cys Phe Thr Asn Pro Leu Pro Ser Asp Tyr Asp Phe Asp  
370 375 380  
Ile Asn Asp Phe Leu Thr Asp Asp Ala Asn His Ala Ala Tyr Asp Ile  
385 390 395 400  
Val Ala Ala Ser Asn Tyr Ala Ala Ala Asp Arg Glu Leu Asp Leu Glu  
405 410 415  
Ile His Asp Pro Glu Asn Gln Ile Pro Ser Arg His Ser Ile Gln Gln  
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Pro Gln Ser Gly Ala Ser Ser His Gly Cys Asp Asp Gly Gly Ile Ala  
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Val Gly Val  
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<211> 349  
<212> PRT  
<213> Aspergillus nidulans

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35 40 45  
Lys Pro Glu Glu Lys Lys Pro Ala Lys Lys Arg Lys Ser Trp Gly Gln  
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Glu Leu Pro Val Pro Lys Thr Asn Leu Pro Pro Arg Lys Arg Ala Lys  
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Arg Ala Ala Ala Gln Thr Ser Arg Glu Arg Lys Arg Leu Glu Met Glu  
100 105 110  
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115 120 125  
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130 135 140  
Gln Val Ala Gln Leu Ser Ala Glu Val Arg Gly Ser Arg His Ser Thr  
145 150 155 160  
Pro Thr Ser Ser Ser Pro Ala Ser Val Ser Pro Thr Leu Thr Pro Thr  
165 170 175  
Leu Phe Lys Gln Glu Gly Asp Glu Val Pro Leu Asp Arg Ile Pro Phe  
180 185 190  
Pro Thr Pro Ser Val Thr Asp Tyr Ser Pro Thr Leu Lys Pro Ser Ser  
195 200 205  
Leu Ala Glu Ser Pro Asp Leu Thr Gln His Pro Ala Val Ser Val Gly  
210 215 220  
Gly Leu Glu Gly Asp Glu Ser Ala Leu Thr Leu Phe Asp Leu Gly Ala  
225 230 235 240  
Ser Ile Lys His Glu Pro Thr His Asp Leu Thr Ala Pro Leu Ser Asp  
245 250 255  
Asp Asp Phe Arg Arg Leu Phe Asn Gly Asp Ser Ser Leu Glu Ser Asp  
260 265 270  
Ser Ser Leu Leu Glu Asp Gly Phe Ala Phe Asp Val Leu Asp Ser Gly  
275 280 285  
Asp Leu Ser Ala Phe Pro Phe Asp Ser Met Val Asp Phe Asp Thr Glu  
290 295 300  
Pro Val Thr Leu Glu Asp Leu Glu Gln Thr Asn Gly Leu Ser Asp Ser  
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<212> DNA

<213> Aspergillus nidulans

<400> 7

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<211> 130

<212> PRT

<213> Aspergillus nidulans

<400> 8

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								20			25		30		
Trp	Asp	Cys	Gln	Ser	Ser	Gln	Ala	Val	Val	Glu	Phe	Val	Arg	Arg	Gly
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Ile	Ala	Ala	Lys	Gln	Asp	Leu	Tyr	Arg	Ile	Cys	Glu	Asn	Met	Met	Asp
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Asn	Cys	Leu	Ala	Ser	Asn	Ser	Glu	Thr	Gly	Gly	Val	Gly	Cys	Asp	Asn
								65			70		75		80
Met	Thr	Met	Val	Ile	Ile	Gly	Leu	Leu	Asn	Gly	Lys	Thr	Lys	Glu	Glu
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Trp	Tyr	Asn	Gln	Ile	Ala	Glu	Arg	Val	Ala	Asn	Gly	Asp	Gly	Pro	Cys
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Ala	Pro	Pro	Glu	Tyr	Gly	Lys	Ser	Leu	Glu	Glu	Pro	Thr	Ala	Ser	Asn
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<210> 9

<211> 1824

<212> DNA

<213> Trichoderma reesei

<400> 9

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acaagatctg	cgagaacatg	atggacaact	gcctgc	caactc	acgggtgg	1080
tcgctgcga	caacatgacc	atggcatca	tcggcttc	gacggca	accaagg	1140
agtgttatg	cgaaattg	aagagagtgg	ccaacgg	cggccctg	gcccccc	1200
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<210> 10

<211> 438

<212> PRT

<213> Trichoderma reesei

<400> 10

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Arg	Ile	Ser	Met	Glu	Asp	Ala	His	Thr	Ala	Glu	Leu	Asn	Leu	Pro	Pro
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Asn	Ile	His	Asn	Ile	Val	Phe	Lys	Gln	Glu	Ser	Phe	Lys	Ser	Gly	Asp
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Tyr	Ala	Gln	Gly	Leu	Lys	Asp	Gly	Phe	Leu	Ala	Thr	Asp	Arg	Ala	Ile
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Val	Thr	Leu	Ile	Ala	Gly	Asn	Lys	Leu	Tyr	Val	Ala	Asn	Ala	Gly	Asp
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Ser	Arg	Ser	Val	Leu	Gly	Ile	Lys	Gly	Arg	Ala	Lys	Pro	Leu	Ser	Asn

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Arg Ala Ile Gly Asp Phe Glu Phe Lys Lys Ser Ala Glu Leu Ser Pro			
195	200	205	
Glu Asn Gln Ile Val Thr Ala Phe Pro Asp Val Glu Val His Glu Leu			
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Thr Glu Glu Asp Glu Phe Leu Val Ile Ala Cys Asp Gly Ile Trp Asp			
225	230	235	240
Cys Gln Ser Ser Gln Ala Val Val Glu Phe Val Arg Arg Gly Ile Ala			
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Ala Lys Gln Asp Leu Asp Lys Ile Cys Glu Asn Met Met Asp Asn Cys			
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Leu Ala Ser Asn Ser Glu Thr Gly Gly Val Gly Cys Asp Asn Met Thr			
275	280	285	
Met Val Ile Ile Gly Phe Leu His Gly Lys Thr Lys Glu Glu Trp Tyr			
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Asp Glu Ile Ala Lys Arg Val Ala Asn Gly Asp Gly Pro Cys Ala Pro			
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Pro Glu Tyr Ala Glu Phe Arg Gly Pro Gly Val His His Asn Tyr Glu			
325	330	335	
Asp Ser Asp Ser Gly Tyr Asp Val Asp Ala Asp Ser Gly Gly Lys Phe			
340	345	350	
Ser Leu Ala Gly Ser Arg Gly Arg Ile Ile Phe Leu Gly Asp Gly Thr			
355	360	365	
Glu Val Leu Thr Gly Ser Asp Asp Thr Glu Met Phe Asp Asn Ala Asp			
370	375	380	
Glu Asp Lys Asp Leu Ala Ser Gln Val Pro Lys Ser Ser Gly Lys Thr			
385	390	395	400
Asp Ala Lys Glu Glu Thr Glu Ala Lys Pro Ala Pro Glu Ala Glu Ser			
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<212> DNA  
<213> Aspergillus nidulans

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<210> 12  
<211> 504  
<212> PRT  
<213> Aspergillus nidulans

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Asp	Ala	Gly	Leu	Ala	Leu	Thr	Arg	Thr	Ala	Ser	Asn	Glu	Val	Phe	Glu
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Asn	Val	Ile	Arg	Cys	Tyr	Cys	Arg	Glu	Gln	Ala	Lys	Gly	Phe	Phe	Tyr
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Gly	Leu	Cys	Lys	Lys	Leu	Glu	Asp	Asn	Gln	Ser	Ser	Phe	Arg	Ala	Thr
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Thr	Ala	His	Ala	Ala	Gly	Thr	Pro	Gly	Gly	Leu	Pro	Asn	Cys	Leu	
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Glu	Ser	Ser	Glu	Pro	Ala	Val	Val	Asp	Pro	Gln	Thr	Asn	Arg	Arg	Ala
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 Pro Phe Phe Trp Asn Pro Ser Asp Arg Leu Ser Phe Leu Cys Asp Val  
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 Ser Asp His Phe Glu Phe Glu Pro Arg Asp Pro Pro Ser Asp Ala Leu  
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 <212> DNA  
 <213> Trichoderma reesei

<400> 13

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<211> 1232

<212> PRT

<213> Trichoderma reesei

<400> 14

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Gln Ile Arg Ile His Ser Gln Arg Gly Asp Ala Pro Leu Asp Lys Val  
35 40 45  
Ala Asp Asp Ala Asn Thr Arg Trp Tyr Ala Thr His Ala Ala Pro Asp  
50 55 60  
Val His Pro Glu Ala Lys Phe Asp Thr Val Asn Arg Lys Gln Lys Gln  
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Gln Ser Thr Ala Ser Pro Gln Gln His Gln Lys Tyr Arg Arg Ala Pro  
85 90 95  
Tyr Asp Tyr Ala Ser Lys Asp Lys Ala Gln Asn Arg Tyr Ala Gln His  
100 105 110  
Pro Ile Arg Glu Ser Glu Lys Pro Asn Tyr Val Lys Val Pro Asn Asp  
115 120 125  
Ala Ser Ala Leu Ala Thr Leu Ala Pro Ala Gln Pro Val Arg Ala Pro  
130 135 140  
His Thr Ser Arg His His Trp Pro Ser Ser Ala Ala Ser Gly Leu  
145 150 155 160  
Ala Ser Pro His Asn Ala Arg Ser Leu Glu Asp Trp Glu Val Glu Asp  
165 170 175  
Phe Val Leu Leu Ala Thr Val Asp Gly Asp Leu Tyr Ala Ser Asp Arg  
180 185 190  
Lys Thr Gly Arg His Leu Trp His Leu Glu Val Asp Gln Pro Val Val  
195 200 205  
Glu Thr Lys His Tyr Arg Thr Asn Asn Ser Val Leu Asp Asp Asp Tyr  
210 215 220  
Arg Pro Val Asp His Tyr Ile Trp Ala Val Glu Pro Ser Arg Asp Gly  
225 230 235 240  
Gly Leu Tyr Val Trp Ile Pro Asp Ser Gly Ala Gly Leu Val Arg Thr  
245 250 255  
Gly Phe Thr Met Lys His Leu Val Glu Glu Leu Ala Pro Tyr Ala Gly  
260 265 270  
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275 280 285  
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305 310 315 320  
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340 345 350  
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405 410 415  
Val Phe Asp Val Cys Arg Pro Trp Asp Ala Asn Ala Gly Ser Asn Pro

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465	470	475
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485	490	495
Ala Phe Asp Ser Ile Asn Pro Asn Lys Leu Ser Lys Ala Leu Val Gly		
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Thr His Phe Leu Asn Pro Val Lys Ser Thr Gly Tyr His Gln Pro Pro		
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Thr Leu Pro Ala Gly Ala Leu Asp Glu Tyr Tyr Glu Asp Leu Glu Asn		
530	535	540
Ala Ser Asn Asn Ala His Ala Val Thr Asn Thr Val Pro Glu Glu Pro		
545	550	555
Thr Ile Ile Thr Lys Val Lys Ala Leu Pro Gln Ser Ala Ala Asn Ser		
565	570	575
Val Ile Asp Phe Val Ser Asn Pro Ile Leu Ile Ile Phe Leu Ile Gly		
580	585	590
Ser Leu Ile Tyr Asn Glu Lys Lys Leu Arg Arg Ser Tyr His Arg Phe		
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Arg Thr His Gly Thr Ile Lys Asp Val Tyr Pro Phe Phe Val Ile Glu		
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Ser Glu Ala Gly Asp Glu Ser Gly Asp Asp Lys Asp Gly Val Phe Pro		
625	630	635
Ser Ser Pro Ser Pro Arg Ser Gln Pro Gln Asp Gln Asn Ala Glu Asp		
645	650	655
His Leu Ser Arg His Lys Val Glu Arg Asn Ala Gly Asp Gln Asp Lys		
660	665	670
Val Lys Asp Asn Arg Ser Leu His Asp Val Ser Asp Thr Leu Glu Pro		
675	680	685
Ser Asn Lys Thr Val Glu Lys Thr Ala Asp Val Val Lys Gln Val Asp		
690	695	700
Val Ala Gly Pro Asp Ala Pro Ser Thr Asp Ser Asn Gly Ala Ala Pro		
705	710	715
Glu Lys Lys Lys Ala His Arg Gly Arg Arg Gly Gly Val Lys His		
725	730	735
Arg Lys Gly Arg Pro Thr Asp Gly Ser Gln Ser His Glu Asn Asp Pro		
740	745	750
Ala Leu Thr Thr Val Asp Glu Ala Val Ser Asn Ala Lys Lys Leu Gly		
755	760	765
Asp Arg Pro Ser Leu Glu Pro Asp Val Met Thr Ile Tyr Asn Asp Met		
770	775	780
Gln Ala Val Thr Gly Ser Val Ile Ser Met Gly Asn Ile Glu Val Asp		
785	790	795
Thr Asp Val Glu Leu Gly Met Gly Ser Asn Gly Thr Val Val Phe Ala		
805	810	815
Gly Arg Phe Asp Gly Arg Asp Val Ala Val Lys Arg Met Thr Ile Gln		
820	825	830
Phe Tyr Asp Ile Ala Thr Arg Glu Thr Lys Leu Leu Arg Glu Ser Asp		
835	840	845
Asp His Pro Asn Val Ile Arg Tyr Tyr Ser Gln Val Gln Arg Gly Asp		
850	855	860
Phe Leu Tyr Ile Ala Leu Glu Arg Cys Ala Ala Ser Leu Ala Asp Val		
865	870	875
		880

Ile Glu Lys Pro Tyr Ala Phe Gly Glu Leu Ala Lys Ala Gly Gln Lys  
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 Asp Leu Pro Gly Val Leu Tyr Gln Ile Thr Asn Gly Ile Ser His Leu  
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 His Ser Leu Arg Ile Val His Arg Asp Leu Lys Pro Gln Asn Ile Leu  
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 Val Asn Leu Asp Lys Asp Gly Arg Pro Arg Leu Leu Val Ser Asp Phe  
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 Gly Leu Cys Lys Lys Leu Glu Asp Arg Gln Ser Ser Phe Gly Ala Thr  
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 Thr Gly Arg Ala Ala Gly Thr Ser Gly Trp Arg Ala Pro Glu Leu Leu  
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 Leu Asp Asp Asp Gly Gln Asn Pro Ala Ala Ile Asp Ser Ser Thr His  
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 Ser Gly Ser His Thr Ile Leu Val Gly Asp Pro Asn Ser Leu Ser Asn  
                   995                  1000                  1005  
 Gly Gly Arg Ala Thr Arg Ala Ile Asp Ile Phe Ser Leu Gly Leu Val  
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 Phe Phe Tyr Val Leu Thr Asn Gly Ser His Pro Phe Asp Cys Gly Asp  
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 Arg Tyr Met Arg Glu Val Asn Ile Arg Lys Gly Asn Tyr Asn Leu Asp  
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 Pro Leu Asp Ala Leu Gly Asp Phe Ala Tyr Glu Ala Lys Asp Leu Ile  
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 Ala Ser Met Leu Gln Ala Ser Pro Lys Ala Arg Pro Asp Ser Arg Glu  
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 Val Met Ala His Pro Phe Phe Trp Ser Pro Lys Lys Arg Leu Ala Phe  
                   1090                  1095                  1100  
 Leu Cys Asp Val Ser Asp Ser Leu Glu Lys Glu Val Arg Asp Pro Pro  
                   1105                  1110                  1115                  1120  
 Pro Ala Leu Val Glu Leu Glu Arg His Ala Pro Glu Val Ile Lys Gly  
                   1125                  1130                  1135  
 Asp Phe Leu Lys Val Leu Thr Arg Asp Phe Val Glu Ser Leu Gly Lys  
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 Gln Arg Lys Tyr Thr Gly Asn Lys Leu Leu Asp Leu Leu Arg Ala Leu  
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 Arg Asn Lys Arg Asn His Tyr Glu Asp Met Ser Asp Ser Leu Lys Arg  
                   1170                  1175                  1180  
 Ser Val Gly Ser Leu Pro Asp Gly Tyr Leu Ala Tyr Trp Thr Val Lys  
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 Phe Pro Met Leu Leu Leu Thr Cys Trp Asn Val Val Tyr Asn Leu Glu  
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attgatccac aattaaagat caatcacctg tcacgcttga aatgatggaa gaagcattct	300
ctccagtcga ctccctcgcc ggctccccga cgccctgagtt gccattgttg acagtgtccc	360

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agaagctgt	gaagaagaga	aagtcatggg	gccaggaatt	gccagtccc	aagactaact	480
tgcccccaag	gaaacggcc	aagactgaag	atgagaaaaga	gcaacgtcg	atcgagcgcg	540
ttcttcgcaa	tcgtgcggca	gcacaacat	cacgcgagcg	caagaggctc	gaaatggaga	600
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cccagatgga	agctgagaac	aatcgcttaa	accaacaagt	cgctcaacta	tctgctgagg	720
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taactcctac	cctatttaaa	caagaacgCG	acgaaatccc	tcttgaacgg	attcccttcc	840
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ccgacgtgac	acaacatcct	gcagcggtgt	tgtgcgacct	gcagtgtccg	tcgctggact	960
cgaaggagaa	ggaagtgcCC	tctctcttt	tgacgtcgGC	teaaaccctg	aacctcacgc	1020
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tgacgtctct	ccttatggc	aagcatagtt	gaggttccgg	ctgtaaatta	tcataaaatcc	1620
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Ala	Ser	Pro	Val	Met	Thr	Arg	Trp	Pro	Val	Phe	Leu	Met	Met	Glu	Glu	
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Ala	Phe	Ser	Pro	Val	Asp	Ser	Leu	Ala	Gly	Ser	Pro	Thr	Pro	Glu	Leu	
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Pro	Leu	Leu	Thr	Val	Ser	Pro	Ala	Asp	Thr	Ser	Leu	Asp	Asp	Ser	Ser	
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Val	Gln	Ala	Gly	Glu	Thr	Lys	Ala	Glu	Glu	Lys	Lys	Pro	Val	Lys	Lys	
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Pro	Arg	Lys	Arg	Ala	Lys	Thr	Glu	Asp	Glu	Lys	Glu	Gln	Arg	Arg	Ile	
								115			120			125		
Glu	Arg	Val	Leu	Arg	Asn	Arg	Ala	Ala	Ala	Gln	Thr	Ser	Arg	Glu	Arg	
								130			135			140		
Lys	Arg	Leu	Glu	Met	Glu	Lys	Leu	Glu	Asn	Glu	Lys	Ile	Gln	Met	Glu	
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Gln	Gln	Asn	Gln	Phe	Leu	Leu	Gln	Arg	Leu	Ser	Gln	Met	Glu	Ala	Glu	
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Asn	Asn	Arg	Leu	Asn	Gln	Gln	Val	Ala	Gln	Leu	Ser	Ala	Glu	Val	Arg	
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Gly	Ser	Arg	Gly	Asn	Thr	Pro	Lys	Pro	Gly	Ser	Pro	Val	Ser	Ala	Ser	
								195			200			205		
Pro	Thr	Leu	Thr	Pro	Thr	Leu	Phe	Lys	Gln	Glu	Arg	Asp	Glu	Ile	Pro	
								210			215			220		

Leu Glu Arg Ile Pro Phe Pro Thr Pro Ser Ile Thr Asp Tyr Ser Pro  
 225 230 235 240  
 Thr Leu Arg Pro Ser Thr Leu Ala Glu Ser Ser Asp Val Thr Gln His  
     245 250 255  
 Pro Ala Val Ser Val Ala Gly Leu Glu Gly Glu Gly Ser Ala Leu Ser  
     260 265 270  
 Leu Phe Asp Val Gly Ser Asn Pro Glu Pro His Ala Ala Asp Asp Leu  
     275 280 285  
 Ala Ala Pro Leu Ser Asp Asp Asp Phe His Arg Leu Phe Asn Val Asp  
     290 295 300  
 Ser Pro Val Gly Ser Asp Ser Ser Val Leu Glu Asp Gly Phe Ala Phe  
     305 310 315 320  
 Asp Val Leu Asp Gly Gly Asp Leu Ser Ala Phe Pro Phe Asp Ser Met  
     325 330 335  
 Val Asp Phe Asp Pro Glu Ser Val Gly Phe Glu Gly Ile Glu Pro Pro  
     340 345 350  
 His Gly Leu Pro Asp Glu Thr Ser Arg Gln Thr Ser Ser Val Gln Pro  
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 Gly Cys  
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   20           25           30  
 Asp Asp Ser Ser Val Gln Ala Gly Glu Thr Lys Ala Glu Glu Lys Lys  
   35           40           45  
 Pro Val Lys Lys Arg Lys Ser Trp Gly Gln Glu Leu Pro Val Pro Lys

50	55	60
Thr Asn Leu Pro Pro Arg	Lys Arg Ala Lys	Thr Glu Asp Glu Lys Glu
65	70	75
Gln Arg Arg Ile Glu Arg Val	Leu Arg Asn Arg Ala Ala	Ala Gln Thr
85	90	95
Ser Arg Glu Arg Lys Arg	Leu Glu Met Glu Lys	Leu Glu Asn Glu Lys
100	105	110
Ile Gln Met Glu Gln Gln Asn	Gln Phe Leu Leu Gln Arg	Leu Ser Gln
115	120	125
Met Glu Ala Glu Asn Asn Arg	Leu Asn Gln Gln Val	Ala Gln Leu Ser
130	135	140
Ala Glu Val Arg Gly Ser Arg Gly Asn Thr	Pro Lys Pro Gly Ser Pro	
145	150	155
Val Ser Ala Ser Pro Thr	Leu Thr Pro Thr	Leu Phe Lys Gln Glu Arg
165	170	175
Asp Glu Ile Pro Leu Glu Arg Ile	Pro Phe Pro Thr Pro	Ser Ile Thr
180	185	190
Asp Tyr Ser Pro Thr Leu Arg Pro	Ser Thr Leu Ala Glu	Ser Ser Asp
195	200	205
Val Thr Gln His Pro Ala Val	Ser Val Ala Gly	Leu Glu Gly Glu Gly
210	215	220
Ser Ala Leu Ser Leu Phe Asp Val	Gly Ser Asn Pro Glu Pro His	Ala
225	230	235
Ala Asp Asp Leu Ala Ala Pro	Leu Ser Asp Asp Asp Phe His	Arg Leu
245	250	255
Phe Asn Val Asp Ser Pro Val	Gly Ser Asp Ser Ser Val	Leu Glu Asp
260	265	270
Gly Phe Ala Phe Asp Val Leu Asp	Gly Gly Asp Leu Ser Ala Phe Pro	
275	280	285
Phe Asp Ser Met Val Asp Phe Asp Pro	Glu Ser Val Gly Phe Glu Gly	
290	295	300
Ile Glu Pro Pro His Gly Leu Pro	Asp Glu Thr Ser Arg Gln Thr	Ser
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36

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<210> 52

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 20 25 30  
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 35 40 45  
 Cys Ser Leu Leu Glu Asn Leu Leu Asn Ser Val Asn Leu Glu Lys Leu  
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